

Insertion of a subretinal prosthesis system for retinitis pigmentosa

Interventional procedures guidance

Published: 16 December 2015

www.nice.org.uk/guidance/ipg537

Your responsibility

This guidance represents the view of NICE, arrived at after careful consideration of the evidence available. When exercising their judgement, healthcare professionals are expected to take this guidance fully into account, and specifically any special arrangements relating to the introduction of new interventional procedures. The guidance does not override the individual responsibility of healthcare professionals to make decisions appropriate to the circumstances of the individual patient, in consultation with the patient and/or guardian or carer.

All problems (adverse events) related to a medicine or medical device used for treatment or in a procedure should be reported to the Medicines and Healthcare products Regulatory Agency using the [Yellow Card Scheme](#).

Commissioners and/or providers have a responsibility to implement the guidance, in their local context, in light of their duties to have due regard to the need to eliminate unlawful

discrimination, advance equality of opportunity, and foster good relations. Nothing in this guidance should be interpreted in a way that would be inconsistent with compliance with those duties. Providers should ensure that governance structures are in place to review, authorise and monitor the introduction of new devices and procedures.

Commissioners and providers have a responsibility to promote an environmentally sustainable health and care system and should assess and reduce the environmental impact of implementing NICE recommendations wherever possible.

1 Recommendations

- 1.1 Current evidence on the safety and efficacy of insertion of a subretinal prosthesis system for retinitis pigmentosa is limited in quality and quantity. Therefore, this procedure should only be used in the context of research.
- 1.2 NICE encourages further research on this procedure. Outcomes should include the impact on quality of life and activities of day-to-day living, and durability of implants. NICE may update the guidance on publication of further evidence.

2 Indications and current treatments

- 2.1 Retinitis pigmentosa is the encompassing term for a group of degenerative eye conditions that cause progressive loss of retinal photoreceptors. The disease is often inherited. Patients initially experience ring scotoma and night vision problems which, in most cases, slowly progress and lead to the loss of all peripheral vision. Central vision is usually preserved until late stages of the disease, but can be lost earlier with severe disease.
- 2.2 Conservative treatments are aimed at early identification and treatment of complications such as cataract or macular oedema. Some newer treatments aim to slow the progression of the condition. Surgical treatments are being developed, including epiretinal and subretinal prostheses, as well as optic nerve implants to restore basic sight.

3 The procedure

- 3.1 Retinitis pigmentosa causes loss of retinal photoreceptors but inner retinal cells (ganglion and bipolar cells) remain intact. Subretinal prosthesis systems aim to restore perception of light, movement, and shapes by surgically implanting a microchip behind the retina. The microchip mimics the function of damaged outer retinal photoreceptors by absorbing light and converting it into retinotopically correct electrical pulses that stimulate the overlying bipolar cell layer. The bipolar cells propagate the signal to downstream retinal cells, which send visual information to the brain.
- 3.2 Implantation of the microchip is done with the patient under general anaesthesia. A vitrectomy is performed and the microchip is implanted underneath the macula using a transscleral, then subretinal approach. The microchip connects to a thin cable that exits the eye at the equator, through the choroid and sclera, and runs under the skin to a power source which is fixed to bone in the retroauricular region. This, in turn, connects to an external power source/control unit via a removable, surface mounted induction loop.

4 Efficacy

This section describes efficacy outcomes from the published literature that the committee considered as part of the evidence about this procedure. For more detailed information on the evidence, see the [interventional procedure overview](#).

- 4.1 In a case series of 6 patients, improvements in visual acuity (measured by the smallest Early Treatment of Diabetic Retinopathy study [ETDRS] letters that could be read) were reported in 3 patients. Visual acuity improved in 1 patient from a Snellen equivalent of 20/800 before the procedure to 20/200 at 6-month follow-up. In the second patient, visual acuity improved from 20/1,600 before the procedure to 20/400 at 6 month follow-up. The third patient had been unable to read ETDRS letters before the procedure but had a visual acuity of 20/1600 at 18 month follow-up.
- 4.2 In a case series of 9 patients, light perception thresholds were considerably

better when prosthesis systems were switched on compared against when they were switched off. All patients were able to perceive light when their prosthesis systems were switched on, at maximum follow-up of 9 months. No further details were provided.

- 4.3 In a case series of 29 patients, patients were asked to indicate whether they saw light when a black screen was briefly illuminated with 1 or 2 flashes of light: a correct response rate of 75% was considered a pass. At 1 month follow-up, 78% of patients passed the test when their prosthesis systems were switched on while 10% of patients passed the test when their prosthesis systems were switched off ($p < 0.05$). At 12 month follow-up, 40% of patients passed the test when their prosthesis systems were switched on while no patients passed the test when their prosthesis systems were switched off ($p < 0.05$).
- 4.4 In the case series of 29 patients, patients were asked to indicate the direction (up, down, left or right) of the pointed end of a white wedge on a black screen: a correct response rate of 62.5% was considered a pass. At 1 month follow-up, 38% of patients passed the test when their prosthesis systems were switched on while no patients passed the test when their prosthesis systems were switched off ($p < 0.05$). At 12 month follow-up, 18% of patients passed the test when their prosthesis systems were switched on while no patients passed the test when their prosthesis systems were switched off (not significant).
- 4.5 In the case series of 9 patients, patients were asked to indicate the direction (up, down, left or right) of the pointed end of a white wedge on a black screen. Seven patients correctly indicated the direction in which the wedge was pointing when their prosthesis systems were switched on, at maximum follow-up of 9 months.
- 4.6 In the case series of 29 patients, patients were asked to indicate the direction of a white polka dot pattern that moved across a black screen: a correct response rate of 62.5% was considered a pass. At 1 month follow-up, 14% of patients passed the test when their prosthesis systems were switched on while no patients passed the test when their prosthesis systems were switched off (not significant). At 12 month follow-up, none of the patients passed the test when their prosthesis systems were switched on or off (not significant).
- 4.7 In the case series of 9 patients, patients were asked to count, locate and identify

4 of 6 possible geometric shapes that were placed on a black table cloth. The mean number of shapes counted was 2.8 when prosthesis systems were switched on, compared against 0.5 when prosthesis systems were switched off, at maximum follow-up of 9 months ($p=0.012$). The mean number of shapes located was 2.2 when prosthesis systems were switched on, compared against 0.5 when prosthesis systems were switched off ($p=0.012$). The mean number of shapes correctly identified was 1 when prosthesis systems were switched on, compared against 0.1 when prosthesis systems were switched off ($p=0.018$).

- 4.8 Specialist advisers listed key efficacy outcomes as improvement of visual function (recognition and discrimination of words or objects, as well as perception of light, movement or direction), performance in spatial or motor tasks (including activities of daily living), and improved quality of life.

5 Safety

This section describes safety outcomes from the published literature that the committee considered as part of the evidence about this procedure. For more detailed information on the evidence, see the [interventional procedure overview](#).

- 5.1 In a case series of 9 patients, 75 adverse events occurred within 1 year of prosthesis implantation. These included:
- 'retinal break' without detachment – 2 cases (3% of adverse events): neither resolved (no further details provided)
 - conjunctival erosions above the external part of the cable, suture erosions through the conjunctiva, or both – 12 cases (16% of adverse events): all resolved without sequelae (no further details provided)
 - conjunctival hyperaemia – 6 cases (8% of adverse events): all resolved without sequelae (no further details provided)
 - retinal vascular leakage and neovascularisation – 10 cases (13% of adverse events): 2 occurred before device implantation. Nine did not resolve. In 1 patient, retinal vascular leakage resulted in damage to eye structures and loss of light perception

- retinal haemorrhage – 7 cases (9% of adverse events): all resolved without sequelae (no further details provided)
- ocular hypertension – 8 cases (11% of adverse events): all resolved without sequelae (no further details provided)
- paraesthesia of the skin (location not specified) – 3 cases (4% of adverse events): all resolved without sequelae (no further details provided)
- epistaxis – 2 cases (3% of adverse events): both resolved without sequelae (no further details provided).

5.2 In the case series of 9 patients a single occurrence of each of the following was reported within 1 year of prosthesis implantation: intraoperative perforation of the choroid, intraoperative contact of the optic nerve head with the implant, postoperative bleeding, contusion of the eyelid and periocular area, mucopurulent conjunctivitis, a peripheral corneal dent, acute iritis, retinal detachment with a retinal break, ocular pain, dizziness, headache, and chronic pain (unspecified location). Intraoperative perforation of the choroid and intraoperative contact of the optic nerve head with the implant both occurred in the same patient and resulted in loss of residual vision in the study eye. All other adverse events resolved without sequelae.

5.3 Device failure was reported in 6 patients in another case series of 9 patients. Cable defects resulted in intraoperative cable breaks in 3 patients: this led to device failure after 3 to 9 months. Corrosion of the periphery of the microchip was reported in 3 patients after 250 days in situ. The chip gradually lost function and the patients opted for explantation of the device.

5.4 In addition to safety outcomes reported in the literature, specialist advisers are asked about anecdotal adverse events (events which they have heard about) and about theoretical adverse events (events which they think might possibly occur, even if they have never done so). For this procedure, specialist advisers listed the following anecdotal adverse events: minor intraoperative subretinal bleeding, and the implant malfunctioning, requiring removal and replacement. They considered that the following were theoretical adverse events: intraocular haemorrhage, glaucoma, photopsia, choroidal neovascularisation, thermal injury to neurons, choroidal or retinal circulation abnormalities, and complications associated with

vitrectomy.

6 Committee comments

- 6.1 The committee noted that insertion of a subretinal prosthesis system for retinitis pigmentosa is intended for patients with end-stage disease who have no useful sight and no other treatment options. It recognised that even minor improvements in vision may help these patients, but it wanted evidence that any changes in metrics of vision result in improvements in quality of life and activities of daily living. These considerations underpinned the specific recommendations about research in [section 1.2](#).
- 6.2 The committee noted that the evidence included studies of different devices, some of which are no longer used. The committee recognised that the technology of subretinal prostheses and related devices is evolving and that further developments may result in substantial changes to outcomes which may influence patient selection in the future.
- 6.3 The committee noted the importance of careful patient selection, including psychological counselling to ensure that patients have realistic expectations. It also noted the need for continued expert care of patients and their subretinal prostheses after the procedure.

7 Further information

Information for patients

NICE has produced [information on this procedure for patients and carers](#). It explains the nature of the procedure and the guidance issued by NICE, and has been written with patient consent in mind.

ISBN: 978-1-4731-1591-0

Endorsing organisation

This guidance has been endorsed by [Healthcare Improvement Scotland](#).